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Page 1 of 51 APPLICATION NO. ATTY. DOCKET NO. AADEMARY 0609.4180002/JAG/JUK 09/785,474 FORM PTO-1449 APPLICANT TANZI et al. INFORMATION DISCLOSURE STATEMENT GROUP FILING DATE February 20, 2001 1647 U.S. PATENT DOCUMENTS EXAMINER NAME CLASS SUB-FILING DATE DOCUMENT DATE INITIAL CLASS NUMBER 04/29/1981 435 68 AA1 4,411,993 10/25/1983 Gillis, S. Frackleton, Jr., A.R. et 935 92 12/13/1982 AB1 4,543,439 09/24/1985 12/14/1981 AC1 Re 32,011 10/22/1985 Zimmerman, T.S. et al. 260 112 B 435 172.3 11/23/1981 04/01/1986 Palmiter, R.D. et al. AD1 4,579,821 6 02/07/1986 435 AE1 4,683,195 07/28/1987 Mullis, K.B. et al. 07/28/1987 Mullis, K.B. 435 91 10/25/1985 4,683,202 AF1 Leder, P. et al. 800 06/22/1984 04/12/1988 AG1 4,736,866 435 235 05/25/1984 AH1 4,786,600 11/22/1988 Kramer, F.R. et al. 172.3 12/17/1986 AI1 4,800,159 01/24/1989 Mullis, K.B. et al. 435 435 6 12/05/1985 Duck, P. et al. AJ1 4,876,187 10/24/1989 435 6 Duck, P. et al. 04/29/1988 5,011,769 04/30/1991 AK1 FOREIGN PATENT DOCUMENTS EXAMINER DOCUMENT DATE COUNTRY CLASS SUB-TRANSLATION INITIAL NUMBER EP 0 125 023 11/14/1984 Yes Europe RUN No AL1 Yes EP 0 171 496 02/19/1986 Europe No AM1 A2 WIPO Yes WO 86/01533 03/13/1986 No AN1 Yes EP 0 173 494 05/03/1986 Europe No A01 A2 EP 0 184 187 06/11/1986 Europe Yes AP1 A2 OTHER (Including Author, Title, Date, Pertinent Pages, etc.) Abraham, C.R. et al., "Immunochemical Identification of the Serine Protease Inhibitor  $\alpha_1$ -Antichymotrypsin in the Brain Amyloid Deposits of Alzheimer's ach AR 1 Disease," Cell 52:487-501, Cell Press (1988) Acsadi, G. et al., "Human dystrophin expression in mdx mice after intramuscular AS 1 injection of DNA constructs," Nature 352:815-818, Macmillan Publishers Ltd (1991) Alber, T. and G. Kawasaki, "Nucleotide Sequence of the Triose Phosphate Isomerase Gene of Saccharomyces cerevisiae, " J. Mol. Appl. Genet. 1:419-434, Raven Press ΑT 1 (1982)DATE CONSIDERED EXAMINER

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	Nishimura, Y. et al., "Recombinant Human-Mouse Chimeric Monoclonal Antibody  Specific for Common Acute Lymphocytic Leukemia Antigen," Cancer Res. 47:999-1005,  American Association for Cancer Research (1987)										
	AT	<u>31</u>	Ohno, T. et al., "Gene Therapy for Vascular Smooth Muscle Cell Proliferation  After Arterial Injury," Science 265:781-784, American Association for the Advancement of Science (August 1994)								
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	Russell, D.R. and G.N. Bennett, "Construction and analysis of in vivo activity of E. coli promoter hybrids and promoter mutants that alter the -35 to -10 spacing,"  Gene 20:231-243, Elsevier Biomedical Press (1982)											
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	AS.	<u>50</u>	Sambrook, J. et al., "Using Mutagenesis to Study Proteins," in Molecular Cloning, A Laboratory Manual, Second Edition, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, New York, pp.15.81-15.84, 16.3-16.4, 17.2 (1989)							
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